

Section 1. Identifying Inform	mation	
Given Name (First Name)  Ta-Chiang	2. Surname (Last Name) Liu	3. Date 18-April-2016
4. Are you the corresponding author?	Yes V No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease p	patients promote dysbiosi	is
6. Manuscript Identifying Number (if you k 86907-INS-CMED-1	know it)	
Section 2. The Work Under C	Consideration for Publ	ication
Did you or your institution at any time reco any aspect of the submitted work (includin statistical analysis, etc.)? Are there any relevant conflicts of inter	g but not limited to grants, d	m a third party (government, commercial, private foundation, etc.) for data monitoring board, study design, manuscript preparation,
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Name of Institution/Company  Doris Duke Charitable Foundation	Grant? Personal No	on-Financial Other? Comments
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Do you have any patents, whether plan	nned, pending or issued, b	proadly relevant to the work? Yes Vo



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Dr. Liu reports grant support from Doris Duke Charitable Foundation.

#### **Evaluation and Feedback**



Section 1. Identifying Inform	mation	
1. Given Name (First Name) Bhaskar	2. Surname (Last Name) Gurram	3. Date 07-April-2016
<ul><li>4. Are you the corresponding author?</li><li>5. Manuscript Title</li></ul>	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
Paneth cell defects in Crohn's disease		
6. Manuscript Identifying Number (if you k 86907-INS-CMED-1	(now it)	
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Do you have any patents, whether plan	nned, pending or issued, bro	oadly relevant to the work? Yes Vo



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Dr. Gurram has nothing to disclose.

### **Evaluation and Feedback**



Section 1. Identifying Infor	mation	
Given Name (First Name)  Megan	2. Surname (Last Name) Baldridge	3. Date 06-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	
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Do you have any patents, whether plan	nned, pending or issued, bro	oadly relevant to the work? Yes V No



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Dr. Baldridge has nothing to disclose.

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Section 1. Identifying Inform	mation	(E)
Given Name (First Name) Richard	2. Surname (Last Name) Head	3. Date 11-April-2016
<ul><li>4. Are you the corresponding author?</li><li>5. Manuscript Title</li><li>Paneth cell defects in Crohn's disease</li></ul>	Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
6. Manuscript Identifying Number (if you k 86907-INS-CMED-1	know it)	
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Dr. Head has nothing to disclose.

#### **Evaluation and Feedback**



Section 1. Identifying Infor	mation	
Given Name (First Name)     VY	2. Surname (Last Name) LAM	3. Date 16-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus Stappenbeck and Nita Salzman
5. Manuscript Title Paneth Cell Defects in Crohn's Disease	Patients Promote Dysbiosi	is
6. Manuscript Identifying Number (if you l 86907-INS-CMED-1	know it)	
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Do you have any patents, whether plan	nned, pending or issued, bro	oadly relevant to the work? Yes Vo



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Dr. LAM has nothing to disclose.

### **Evaluation and Feedback**



1. Given Name (First Name) Chengwei	2. Surname (Last Name) Luo	3. Date 11-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	Tradacas S. Stappenseck, Mita H. Suizman
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Dr. Luo has nothing to disclose.
Evaluation and Feedback



1. Given Name (First Name) Yumei	2. Surname (Last Name) Cao	3. Date 07-April-2016
4. Are you the corresponding author?	Yes 🗸 No	Corresponding Author's Name
5. Manuscript Title Paneth cell defects in Crohn's disease	e patients promote dysbiosis	Thaddeus S. Stappenbeck, Nita H. Salzman
6. Manuscript Identifying Number (if you 86907-INS-CMED-1	know it)	
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Dr. Cao has nothing to disclose.

#### **Evaluation and Feedback**



Section 1. Identifying Inform	nation	
Given Name (First Name)  pippa	2. Surname (Last Name) simpson	3. Date 07-April-2016
4. Are you the corresponding author?	☐ Yes   ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease p	patients promote dysbiosis	
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Do you have any patents, whether plan	ned, pending or issued, bro	padly relevant to the work? Yes Vo



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Section 1. Identifying Infor	mation	
Given Name (First Name)     Michael	2. Surname (Last Name) Hayward	3. Date 18-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	
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Dr. Hayward has nothing to disclose.

### **Evaluation and Feedback**



Section 1. Identifying Infor	mation	
Given Name (First Name)  Mary	2. Surname (Last Name) Holtz	3. Date 11-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	
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Dr. Holtz has nothing to disclose.

## **Evaluation and Feedback**



Section 1. Identifying Inform	mation	STEEL STATE OF STATE OF STREET STATE OF STREET STATE OF STREET
Given Name (First Name) PAVLOS	2. Surname (Last Name) BOUSOUNIS	3. Date 12-April-2016
<ul><li>4. Are you the corresponding author?</li><li>5. Manuscript Title</li><li>Paneth cell defects in Crohn's disease p</li></ul>	Yes No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
6. Manuscript Identifying Number (if you k 86907-INS-CMED-1	now it)	
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Mr. BOUSOUNIS has nothing to disclose.

## **Evaluation and Feedback**



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5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis		y
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Section 5. Relationships not covered above
Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?
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Dr. Noe has nothing to disclose.

## **Evaluation and Feedback**



Section 1. Identifying Inform	ation	
Given Name (First Name)  Diana	2. Surname (Last Name) Lerner	3. Date 06-April-2016
<ul><li>4. Are you the corresponding author?</li><li>5. Manuscript Title</li></ul>	Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
Paneth cell defects in Crohn's disease pa	atients promote dysbiosis	
6. Manuscript Identifying Number (if you kn 86907-INS-CMED-1	ow it)	
Section 2. The Work Under Co	onsideration for Public	ation
Did you or your institution <b>at any time</b> receivany aspect of the submitted work (including statistical analysis, etc.)?	ve payment or services from a but not limited to grants, dat	a third party (government, commercial, private foundation, etc.) for a monitoring board, study design, manuscript preparation,
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Section 4. Intellectual Property	y Patents & Copyrigh	**
Do you have any patents, whether planne	2002	



Section 5. Relationships not covered above
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Dr. Lerner has nothing to disclose.

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Section 1. Identifying Inform	nation	
Given Name (First Name)  Jose	2. Surname (Last Name) Cabrera	3. Date 11-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
<ol><li>Manuscript Title</li><li>Paneth cell defects in Crohn's disease p</li></ol>	patients promote dysbiosis	
6. Manuscript Identifying Number (if you kr 86907-INS-CMED-1	now it)	
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Section 4. Intellectual Proper	ty Patents & Copyrigl	
Do you have any patents, whether plann		Company of the State of King of the State of
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Section 5. Relationships not covered above
Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?
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Dr. Cabrera has nothing to disclose.

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1. Given Name (First Name)	2. Surname (Last Name)		
Vincent	Biank	3. Date 07-April-2016	
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name	
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	Thaddeus S. Stappenbeck, Nita H. Salzman	
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Dr. Biank has nothing to disclose.

## **Evaluation and Feedback**



Section 1. Identifying Inform	nation	
Given Name (First Name)     Michael	2. Surname (Last Name) Stephens	3. Date 11-April-2016
<ul><li>4. Are you the corresponding author?</li><li>5. Manuscript Title</li><li>Paneth cell defects in Crohn's disease parts.</li></ul>	Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
6. Manuscript Identifying Number (if you kn 86907-INS-CMED-1		
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Section 4. Intellectual Propert	y Patents & Copyrigl	nts
Do you have any patents, whether plann	ed, pending or issued, bro	adly relevant to the work? Yes No



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Dr. Stephens has nothing to disclose.

#### **Evaluation and Feedback**



1. Given Name (First Name) Curtis	2. Surname (Last Name) Huttenhower	3. Date 18-April-2018
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	
6. Manuscript Identifying Number (if you 86907-INS-CMED-1	know it)	
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I am a scientific advisor for Seres Therapeutics, Evelo Therapeutics, and Microbiome Insights; to the best of my knowledge none of them involve work relevant to this publication.
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Dr. Huttenhower reports and I am a scientific advisor for Seres Therapeutics, Evelo Therapeutics, and Microbiome Insights; to the best of my knowledge none of them involve work relevant to this publication

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Section 1. Identifying Infor	mation		
Given Name (First Name)  Dermot	2. Surname (Last Name) McGovern		3. Date 11-April-2016
4. Are you the corresponding author?	☐ Yes ✓ No	Corresponding Author's Thaddeus S. Stappenb	
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbios	sis	
6. Manuscript Identifying Number (if you 86907-INS-CMED-1	know it)		
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If yes, please fill out the appropriate in Excess rows can be removed by pressi	formation below. If you h	ave more than one entity p	press the "ADD" button to add a row.
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Do you have any patents, whether planned, pending or issued, broadly relevant to the work? Yes Vo
Section 5. Relationships not covered above
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Dr. McGovern reports grants from NIH, European Union, Helmsley Charitable Trust, during the conduct of the study; personal fees from Merck, personal fees from Jannsen, personal fees from UCB, outside the submitted work; .

### **Evaluation and Feedback**



Section 1. Identifying Inform	nation	
Given Name (First Name)     Ramnik	2. Surname (Last Name) Xavier	3. Date 11-April-2016
4. Are you the corresponding author?	Yes 🗸 No	Corresponding Author's Name Thaddeus S. Stappenbeck, Nita H. Salzman
5. Manuscript Title Paneth cell defects in Crohn's disease p	atients promote dysbiosis	
6. Manuscript Identifying Number (if you kr 86907-INS-CMED-1	now it)	
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Section 4. Intellectual Proper	ty Patents & Copyrig	ıhts
Do you have any patents, whether plan	ned, pending or issued, br	oadly relevant to the work? Yes Vo



Section 5. Relationships not covered above
Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?
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Section 6. Disclosure Statement
Based on the above disclosures, this form will automatically generate a disclosure statement, which will appear in the box below.
Dr. Xavier has nothing to disclose.

### **Evaluation and Feedback**



Section 1. Identifying Infor	mation	
Given Name (First Name)     Thaddeus	2. Surname (Last Name) Stappenbeck	3. Date 11-April-2016
4. Are you the corresponding author?	✓ Yes No	
5. Manuscript Title Paneth cell defects in Crohn's disease	patients promote dysbiosis	
6. Manuscript Identifying Number (if you 86907-INS-CMED-1	know it)	
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Section 5. Relationships not covered above
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Dr. Stappenbeck has nothing to disclose.

### **Evaluation and Feedback**



Section 1.			
Identifying Inform	nation		
Given Name (First Name)     Nita	2. Surname (Last Name) Salzman		3. Date 08-April-2016
4. Are you the corresponding author?	✓ Yes No		
5. Manuscript Title Paneth cell defects in Crohn's disease p	atients promote dysbiosis		
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Section 3. Relevant financial	activities outside the sub	mitted work.	<b>从外国外</b> "是"
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Are there any relevant conflicts of inter	est? Yes 🗸 No		
Section 4. Intellectual Prope	rty Patents & Copyright	s and the same	
Do you have any patents, whether plan	ned, pending or issued, broad	dly relevant to the work?	Yes Vo



Section 5. Relationships not covered above
Are there other relationships or activities that readers could perceive to have influenced, or that give the appearance of potentially influencing, what you wrote in the submitted work?
Yes, the following relationships/conditions/circumstances are present (explain below):  No other relationships/conditions/circumstances that present a potential conflict of interest
At the time of manuscript acceptance, journals will ask authors to confirm and, if necessary, update their disclosure statements. On occasion, journals may ask authors to disclose further information about reported relationships.
Section 6. Disclosure Statement
Based on the above disclosures, this form will automatically generate a disclosure statement, which will appear in the box below.
Dr. Salzman reports grants from Crohn's and Colitis Foundation, during the conduct of the study; .

### **Evaluation and Feedback**

 $Please\ visit\ \underline{http://www.icmje.org/cgi-bin/feedback}\ to\ provide\ feedback\ on\ your\ experience\ with\ completing\ this\ form.$ 

## Paneth Cell Defects in Crohn's Disease Patients Promote Dysbiosis (86907-INS-CMED-1)

The following checklist was used for preparation of the study described in the manuscript. All required items have been met.

STROBE Statement—Checklist of items that should be included in reports of cross-sectional studies

	Item No	Recommendation
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract
		(b) Provide in the abstract an informative and balanced summary of what was done
		and what was found
Introduction		
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported
Objectives	3	State specific objectives, including any prespecified hypotheses
Methods		
Study design	4	Present key elements of study design early in the paper
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment,
C		exposure, follow-up, and data collection
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of
•		participants
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect
		modifiers. Give diagnostic criteria, if applicable
Data sources/	8*	For each variable of interest, give sources of data and details of methods of
measurement		assessment (measurement). Describe comparability of assessment methods if there is
		more than one group
Bias	9	Describe any efforts to address potential sources of bias
Study size	10	Explain how the study size was arrived at
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable,
		describe which groupings were chosen and why
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain how missing data were addressed
		(d) If applicable, describe analytical methods taking account of sampling strategy
		$(\underline{e})$ Describe any sensitivity analyses
Results		
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially
		eligible, examined for eligibility, confirmed eligible, included in the study,
		completing follow-up, and analysed
		(b) Give reasons for non-participation at each stage
		(c) Consider use of a flow diagram
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and
		information on exposures and potential confounders
		(b) Indicate number of participants with missing data for each variable of interest
Outcome data	15*	Report numbers of outcome events or summary measures
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and
		their precision (eg, 95% confidence interval). Make clear which confounders were
		adjusted for and why they were included
		(b) Report category boundaries when continuous variables were categorized

		(c) If relevant, consider translating estimates of relative risk into absolute risk for a
		meaningful time period
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and
		sensitivity analyses
Discussion		
Key results	18	Summarise key results with reference to study objectives
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or
		imprecision. Discuss both direction and magnitude of any potential bias
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations,
		multiplicity of analyses, results from similar studies, and other relevant evidence
Generalisability	21	Discuss the generalisability (external validity) of the study results
Other information		
Funding	22	Give the source of funding and the role of the funders for the present study and, if
		applicable, for the original study on which the present article is based

<sup>\*</sup>Give information separately for exposed and unexposed groups.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.